

PCT WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)



(51) International Patent Classification 7:

A61K 39/39, 39/095, 39/102, 39/106, 39/245, 39/155, 39/15, C12N 15/63, 5/10 // C07K 14/28

(11) International Publication Number:

DOEDE

WO 00/18434

(43) International Publication Date:

6 April 2000 (06.04.00)

(21) International Application Number:

PCT/US99/22520

A1

(22) International Filing Date:

30 September 1999 (30.09.99)

(30) Priority Data:

60/102,430

30 September 1998 (30.09.98) US

(71) Applicants (for all designated States except US): AMER-ICAN CYANAMID COMPANY [US/US]; Five Giralda Farms, Madison, NJ 07940 (US). THE GOVERNMENT OF THE UNITED STATES OF AMERICA as represented by the UNIFORMED SERVICES UNIVERSITY OF THE HEALTH SCIENCES [US/US]; 4301 Jones Bridge Road, Bethesda, MD 20814-4799 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): HOLMES, Randall, K. [US/US]; 23371 Morning Rose Drive, Golden, CO 80401 (US). JOBLING, Michael, G. [GB/US]; 5250 Cherry Creek Drive South, Apt. 18A, Denver, CO 80222 (US). EL-DRIDGE, John, H. [US/US]; 4 Beatrice Cove, Fairport, NY 14450 (US). GREEN, Bruce, A. [US/US]; 49 Northfield Gate, Pittsford, NY 14534 (US). HANCOCK, Gerald, E. [US/US]; 50 Plains Road, Honeoye Falls, NY 14472 (US).

PEEK, Joel, A. [US/US]; 156 Railroad Mills Road, Pittsford, NY 14534 (US).

- (74) Agents: GORDON, Alan, M.; American Home Products Corporation, Patent Law Dept. - 2B2, One Campus Drive, Parsippany, NJ 07054 (US) et al.
- (81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published

With international search report.

Before the expiration of the time limit for amending the claims and to be republished in the event of the receipt of amendments.

(54) Title: MUTANT CHOLERA HOLOTOXIN AS AN ADJUVANT

(57) Abstract

A mutant cholera holotoxin featuring a point mutation at amino acid 29 of the A subunit, wherein the glutamic acid residue is replaced by an amino acid other than aspartic acid, is useful as an adjuvant in an antigenic composition to enhance the immune response in a vertebrate host to a selected antigen from a pathogenic bacterium, virus, fungus or parasite. In a particular embodiment, the amino acid 29 is histidine. The mutant cholera holotoxin may contain at least one additional mutation in the A subunit at a position other than amino acid 29. The antigenic composition may include a second adjuvant in addition to the mutant cholera holotoxin.





FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav	TM	Turkmenistan
BF	Burkina Faso	GR	Greece		Republic of Macedonia	TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand		
CM	Cameroon		Republic of Korea	PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		